

Supporting information

Critical view on the qualification of electronic tongues regarding their performance in the development of peroral drug formulations with bitter ingredients

Denise Steiner¹, Alexander Meyer², L. Isabell Immohr³, Miriam Pein-Hackelbusch^{2,*}

¹Institute of Pharmaceutical Technology and Biopharmaceutics, University of Muenster, Corrensstraße 48, 48149 Muenster, Germany

²Institute for Life Science Technologies (ILT.NRW), Ostwestfalen-Lippe University of Applied Sciences and Arts, Campusallee 12, 32657 Lemgo, Germany

³NextPharma Germany Bidco GmbH, Hildebrandstrasse 12, 37081 Goettingen, Germany

Table S1: Summarized information about References found in between 2014-2023; CPA = change in membrane potential due to adsorption, PCA = principal component analysis, PLS = partial least squares regression, DFA = discriminant function analysis, BATA = brief aversion taste assessment – performed with rodents, NCE = new chemical entity; HME = hot melt extrudates; ODT = orodispersible tablets; ODF = orodispersible films

Year	Applied electronic tongue	Taste-masking evaluation by					Analysed drugs	Formulation	References
		Dedicated sensors	CPA value	(Statistical) analysis of electronic tongue data	Human data	Others			
2014	oASTREE II			PCA			Acetaminophen Ibuprofen Tramadol HCl Sildenafil citrate	Solution	[58]
	oASTREE II			PCA	x		Ebastine	ODT	[132]
	Inset TS-5000Z oASTREE II	SB2BT0		PCA and PLS	x		Propranolol HCl	HME	[98]
	Inset TS-5000Z oASTREE II			PCA			Loperamide HCl Dextromethorphan HBr	Solution	[133]
	oASTREE II			PCA and DFA	x		Sildenafil citrate	Microcapsules	[57]
2015	Inset TS-5000Z			PCA			Diclofenac Diclofenac sodium Diclofenac potassium	ODT	[55]
	oASTREE			PCA and PLS	x		Famotidine Amlodipine besylate	ODT	[89]
	Inset TS-5000Z			PCA	x		Cetirizine HCl	ODT	[93]
2016	ASTREE			PCA			Ketoprofen	HME	[96]
2017	Laboratory prototype potentiometric e-tongue			PCA	x		Cetirizine 2HCl	Powder	[80]
	Laboratory prototype potentiometric e-tongue			PCA	x		Cetirizine 2HCl	ODT, Lyophilisates	[90]
	oASTREE			PCA, model based on e-Tongue Euclidean distance measurements, Ka and Kd of the drug/cyclodextrin-complexes			44 bitter drugs	Solution	[71]
	Inset TS-5000Z			PCA	x		Two NCEs	Solution	[5]
	ASTREE			Bitterness score (<i>calculated with electronic tongue software</i>)			Mefloquine HCl	Liposomes	[73]
	Laboratory prototype potentiometric e-tongue			PCA	x		Cetirizine 2HCl	Microspheres	[81]

Year	Applied electronic tongue	Taste-masking evaluation by					Analysed drugs	Formulation	References
		Dedicated sensors	CPA value	(Statistical) analysis of electronic tongue data	Human data	Others			
2018	Inset TS-5000Z			PCA			Ranitidine HCl	Solution	[134]
	ASTREE II			PCA	x		Traditional chinese herbs	Solution	[135]
	ASTREE II			PCA	x	BATA	Berberine HCl	Minitablets	[100]
	Inset TS-5000Z	SB2AC0 SB2AN0 SB2C00 SB2AE1		PCA		BATA	Isoniazide	HME	[97]
	Laboratory prototype electrochemical impedance spectroscopy-based e-tongue			multidimensional projection techniques such as Sammon's Mapping (SAMMON) and PCA but with Interactive Document Map showing the best results			Praziquantel	Microcapsules	[136]
	Inset SA402B	SB2BT0		PCA			Quinine HCl x 2 H ₂ O	HME	[137]
2019	Inset TS-5000Z	SB2C00 SB2AE1 SB2AC0 SB2AN0		Sensor signal			Chlorpheniramine maleate	Solution	[138]
	ASTREE II			PCA			Rhei radix et rhizome Berberine HCl Scutellaria baicalensis	Powder	[139]
	aASTREE			PCA	x		Nizatidine	Powder	[84]
	Laboratory prototype voltammogramm-based e-tongue			PCA	x		Efavirenz	SNEDDS	[105]
	Laboratory prototype potentiometric e-tongue			PCA	x		Rupatadine fumarate	Powder	[83]
2020	Inset TS-5000Z	SB2BT0 SB2AN0 SB2C00 SB2AE1	x	Sensor signal		for veterinary applications	Amlodipine besylate	Minitablets	[99]
	Inset TS-5000Z	SB2BT0 SB2AN0 SB2C00 SB2AE1	x	PCA			Cefpodoxime proxetil	Solution	[140]
	Inset TS-5000Z	SB2AC0 SB2AN0 SB2C00 SB2AE1		Sensor signal	x	BATA	Isoniazid Rifampicin Pyrazinamide Ethambutol 2HCl	Solution	[106]
	Inset SA402B	SB2BT0 (SB2AN0 SB2C00 SB2AE1)	x	PCA	x		Atomoxetine HCl	Solution	[70]
	Inset SA402B	SB2AC0 SB2AN0	x	<i>bitterness intensity estimated as:</i> <i>bitterness intensity 1 = 1.26 × CPA (AC0),</i> <i>bitterness intensity 2 = 1.80 × CPA (AN0)</i> <i>bitterness scores calculated on bitterness intensities</i> <i>total bitterness = (bitterness score 1 + bitterness score 2)/2</i>			Propranolol HCl	Microbeads	[141]

	Laboratory prototype potentiometric e-tongue		PCA	x		Rupatadine fumarate	ODT	[88]
--	--	--	-----	---	--	---------------------	-----	------

Year	Applied electronic tongue	Taste-masking evaluation by					Analysed drugs	Formulation	References
		Dedicated sensors	CPA value	(Statistical) analysis of electronic tongue data	Human data	Others			
2021	Inset TS-5000Z	SB2BT0	X	PCA			Chlorpheniramine maleate	Nanofibres	[103]
	Inset TS-5000Z	SB2AE1 SB2CT0 SB2AC0 SB2AN0 SB2C00		PCA			Indomethacin Furosemide	ODF	[101]
	Laboratory prototype potentiometric e-tongue			PCA			Ketoprofen lysine	Solution	[142]
	Laboratory prototype electrochemical impedance spectroscopy-based e-tongue			Interactive Document Map (IDMAP)		BATA	Artesunate-Mefloquine Praziquantel, Benznidazole	Tablets	[16]
	αASTREE			Bitterness level (response value R)	x	BATA	Ibuprofen	Emulsion	[56]
	Inset TS-5000Z	SB2C00		Bitterness intensity (single sensor response)			Caffeine	Powder	[143]
	ASTREE			PCA, PLS	x		Levetiracetam	ODT	[91]
	Laboratory prototype taste biosensor (rat cardiomyocytes + microelectrode assay)			Multiple feature extraction and parameter visualization for high-content analysis			Denatonium benzoate Quinine Dephenidol Arbutin	Solution	[144]
2022	ASTREE II			PCA			Levocetirizine HCl	Nanofibres	[104]
	Laboratory prototype potentiometric e-tongue			Supervised LDA			Ranitidine HCl	Solution	[74]
	Laboratory prototype potentiometric e-tongue			PCA	x		Rupatadine fumarate	ODFs	[145]
	cTongue; Shanghai Baosheng, Shanghai, China			PCA			Berberine Phillyrin	Solution	[146]
	Inset SA402B	SB2AE1 SB2CT0 SB2AAE SB2CA0 SB2C00 SB2GL1		Peptides were dissolved in water and added to an 1 mM quinine HCl solution – bitterness intensity reduction activity was calculated based on the Δ sensor responses against quinine HCl as blank control and a particular peptide (LEGSLE) as positive control			Nebulin-derived peptides	Solution	[147]
	Inset TS-5000Z	SB2BT0 SB2AN0 SB2C00 SB2AE1	X	Sensor signal	x		Oseltamivir phosphate	HME	[87]
2023	Laboratory prototype potentiometric e-tongue			PCA			Rosuvastatin calcium	Granules (pellets)	[86]
	ASTREE			Relative distances	x		Azithromycin	Granules (pellets)	[85]
	Inset TS-5000Z	SB2C00 SB2AE1 SB2AN0	X	Sensor signal			Carbamazepine	ODT	[92]

	Insent SA402B	umami, saltiness, sourness, bitterness	x	Random Forest, backpropagation neural network, K-nearest neighbour, support vector machine		uf-GC e-nose	Bear pile powder	Powder	[148]
	Insent TS-5000Z	SB2C00 SB2AN0 SB2BT0 SB2AE1	x	Bitter value	x		Berberine HCl	Solution	[75]
Year	Applied electronic tongue			Taste-masking evaluation by			Analysed drugs	Formulation	References
	Dedicated sensors			CPA value	(Statistical) analysis of electronic tongue data		Human data	Others	
2023	Insent SA402B	SB2AN0	x	Sensor signal	x		Ibuprofen Flurbiprofen Naproxen	Solution	[149]
	Insent SA402B	SB2CA0 SB2C00 SB2AE1 SB2AAE SB2CT0	x	PCA			Lisdexamfetamine	Tablets	[150]
	Insent SA402B	SB2BT0	x	Sensor signal			Diphenhydramine HCl	Granules	[82]
	Insent SA402B	SB2AC0 SB2AN0 SB2C00 SB2AE1		PCA and calculation of bitterness scores			Propranolol HCl	Solution	[72]

Klicken oder tippen Sie hier, um Text einzugeben.